

Sequence Listing

<110> Botstein,David

Desnoyers,Luc

Ferrara,Napoleone

Fong,Sherman

Gao,Wei-Qiang

Goddard,Audrey

Gurney,Austin L.

Pan,James

Roy,Margaret Ann

Stewart,Timothy A.

Tumas,Daniel

Watanabe,Colin K.

Wood,William I.

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| Gln | Arg | Ala | Gly | Ala 140 | Arg | Pro | Gly | Ser | Leu 145 | Pro | Glu | Arg | Lys | Gly 150 |
| Pro | Glu | Leu | Cys | Leu 155 | Glu | Glu | Leu | Asp | Ala 160 | Ala | Ile | Pro | Gly | Ser 165 |
| Arg | Ala | Val | Gly | Gly 170 | Ser | Lys | Ala | Arg | Val 175 | Gln | Ala | Arg | Gln | Val 180 |
| Pro | Pro | Ala | Thr | Ala 185 | Ser | Glu | Trp | Arg | Leu 190 | Ala | Gln | Ala | Gln | Gln 195 |
| Lys | Ile | Arg | Glu | Leu 200 | Ala | Ile | Asn | Ile | Arg 205 | Met | Lys | Glu | Glu | Leu 210 |
| Ile | Gly | Glu | Leu | Val 215 | Arg | Thr | Gly | Lys | Ala 220 | Ala | Gln | Ala | Leu | Asn 225 |
| Arg | Gln | His | Ser | Gln 230 | Arg | Ile | Arg | Glu | Leu 235 | Glu | Gln | Glu | Ala | Glu 240 |
| Gln | Val | Arg | Ala | Glu 245 | Leu | Ser | Glu | Gly | Gln 250 | Arg | Gln | Leu | Arg | Glu 255 |
| Leu | Glu | Gly | Lys | Glu 260 | Leu | Gln | Asp | Ala | Gly 265 | Glu | Arg | Ser | Arg | Leu 270 |
| Gln | Glu | Phe | Arg | Arg 275 | Arg | Val | Ala | Ala | Ala 280 | Gln | Ser | Gln | Val | Gln 285 |
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| Gln | Leu | Met | Arg | Gln 320 | Gln | Gln | Gly | Gln | Leu 325 | Gln | Arg | Arg | Leu | Arg 330 |
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| Ser Gln Ala Leu | Asn Glu Asp Ile Val | Arg Val Ser Ser Arg Leu | | | |
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| Glu His Leu Glu | Lys Glu Leu Ser Glu | Lys Ser Gly Gln Leu Arg | | | |
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| Gln Gly Ser Ala | Gln Ser Gln Gln Gln | Ile Arg Gly Glu Ile Asp | | | |
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| Glu Arg Thr Leu | Phe Gln Leu Asp Glu | Ala Ile Glu Ala Leu Asp | | | |
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| Leu Leu Cys Lys | Tyr Phe Asp Lys Val | Val Thr Leu Arg Glu Glu | | | |
| | 590 | | 595 | | 600 |
| Gln His Gln Gln | Gln Ile Ala Phe Ser | Glu Leu Glu Met Gln Leu | | | |
| | 605 | | 610 | | 615 |
| Glu Glu Gln Gln | Arg Leu Val Tyr Trp | Leu Glu Val Ala Leu Glu | | | |
| | 620 | | 625 | | 630 |
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| Glu His Glu Gln | Asn Met Gln Leu Leu | Leu Gln Gln Ser Arg Asp | | | |
| | 650 | | 655 | | 660 |
| His Leu Gly Glu | Gly Leu Ala Asp Ser | Arg Arg Gln Tyr Glu Ala | | | |
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| Arg Ile Gln Ala | Leu Glu Lys Glu Leu | Gly Arg Tyr Met Trp Ile | | | |
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| | | | | | | | | | | | | | | | |
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| Gln | Ala | Pro | Gly | Asn | Glu | Asp | Glu | Leu | His | Leu | Ala | Pro | Glu | Leu | |
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| Leu | Trp | Leu | Ser | Pro | Leu | Thr | Glu | Gly | Ala | Pro | Arg | Thr | Arg | Glu | |
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 Trp Pro Cys Phe Glu Leu Cys Cys Pro Glu Ser Phe Gly Pro Gln
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Ser Lys Leu Lys Gln Ala Trp Val Ser Gln Gly Gly Gly Ala Lys
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110 115 120
Glu Ala Leu Ala Ala Ala Ala Gln Asp Ala Ile Gly Pro Glu Leu
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Ala Pro Thr Pro Glu Pro Pro Glu Glu Tyr Val Tyr Pro Asp Tyr
140 145 150
Arg Gly Lys Gly Cys Val Asp Glu Ser Gly Phe Val Tyr Ala Ile
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Gly Glu Lys Phe Ala Pro Gly Pro Ser Ala Cys Pro Cys Leu Cys
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Thr Glu Glu Gly Pro Leu Cys Ala Gln Pro Glu Cys Pro Arg Leu
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His Pro Arg Cys Ile His Val Asp Thr Ser Gln Cys Cys Pro Gln
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Cys Lys Glu Arg Lys Asn Tyr Cys Glu Phe Arg Gly Lys Thr Tyr
215 220 225
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Asn Thr Ser Cys Asp Ser Gly Leu Gly Cys Gln Asp Thr Leu Met
50 55 60
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65 70 75
Cys Thr Glu Ala Lys Asp Gln Glu Pro Arg Val Thr Glu His Arg
80 85 90
Met Gly Pro Gly Leu Ser Leu Ile Ser Tyr Thr Phe Val Cys Arg
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Gln Glu Asp Phe Cys Asn Asn Leu Val Asn Ser Leu Pro Leu Trp
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| Leu | Trp | Tyr | Ala | Asp 140 | Trp | Met | Asp | Gly | Gln 145 | Leu | Val | Glu | Lys | Gly 150 |
| Arg | Gln | Asp | Met | Arg 155 | Gln | Leu | Ala | Leu | Arg 160 | Leu | Ala | Ser | Leu | Phe 165 |
| Pro | Ala | Leu | Phe | Ser 170 | Arg | Glu | Asn | Tyr | Gly 175 | Arg | Leu | Arg | Leu | Ile 180 |
| Thr | Ser | Ser | Lys | His 185 | Arg | Cys | Met | Asp | Ser 190 | Ser | Ala | Ala | Phe | Leu 195 |
| Gln | Gly | Leu | Trp | Gln 200 | His | Tyr | His | Pro | Gly 205 | Leu | Pro | Pro | Pro | Asp 210 |
| Val | Ala | Asp | Met | Glu 215 | Phe | Gly | Pro | Pro | Thr 220 | Val | Asn | Asp | Lys | Leu 225 |
| Met | Arg | Phe | Phe | Asp 230 | His | Cys | Glu | Lys | Phe 235 | Leu | Thr | Glu | Val | Glu 240 |
| Lys | Asn | Ala | Thr | Ala 245 | Leu | Tyr | His | Val | Glu 250 | Ala | Phe | Lys | Thr | Gly 255 |
| Pro | Glu | Met | Gln | Asn 260 | Ile | Leu | Lys | Lys | Val 265 | Ala | Ala | Thr | Leu | Gln 270 |
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| Phe | Thr | Cys | Ser | Phe 290 | Asp | Leu | Ala | Ile | Lys 295 | Gly | Val | Lys | Ser | Pro 300 |
| Trp | Cys | Asp | Val | Phe 305 | Asp | Ile | Asp | Asp | Ala 310 | Lys | Val | Leu | Glu | Tyr 315 |
| Leu | Asn | Asp | Leu | Lys 320 | Gln | Tyr | Trp | Lys | Arg 325 | Gly | Tyr | Gly | Tyr | Thr 330 |
| Ile | Asn | Ser | Arg | Ser 335 | Ser | Cys | Thr | Leu | Phe 340 | Gln | Asp | Ile | Phe | Gln 345 |
| His | Leu | Asp | Lys | Ala 350 | Val | Glu | Gln | Lys | Gln 355 | Arg | Ser | Gln | Pro | Ile 360 |
| Ser | Ser | Pro | Val | Ile 365 | Leu | Gln | Phe | Gly | His 370 | Ala | Glu | Thr | Leu | Leu 375 |
| Pro | Leu | Leu | Ser | Leu 380 | Met | Gly | Tyr | Phe | Lys 385 | Asp | Lys | Glu | Pro | Leu 390 |
| Thr | Ala | Tyr | Asn | Tyr 395 | Lys | Lys | Gln | Met | His 400 | Arg | Lys | Phe | Arg | Ser 405 |

100224 "H2E00T"

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 Gln Asp Leu Glu Pro Gln Gln Leu Leu Leu Phe Val Gln Ser Phe
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 Gly Ile Pro Val Ser Ser Met Ser Lys Leu Leu Gln Phe Leu Asp
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| | | | | | | | | | | | | | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Gln | Ala | Val | Ala | His | Asp | Pro | Gln | Thr | Leu | Glu | Gln | Asn | Ile | Met | 110 | 115 | 120 |
| Asp | Lys | Asn | Tyr | Met | Ala | His | Leu | Val | Glu | Val | Gln | His | Glu | Arg | 125 | 130 | 135 |
| Gly | Ala | Ser | Gly | Gly | Gln | Thr | Phe | His | Ser | Leu | Leu | Thr | Ala | Ser | 140 | 145 | 150 |
| Leu | Pro | Pro | Arg | Arg | Asp | Ser | Thr | Glu | Ala | Pro | Lys | Pro | Lys | Ser | 155 | 160 | 165 |
| Ser | Pro | Glu | Gln | Pro | Ile | Gly | Gln | Gly | Arg | Ile | Arg | Val | Gly | Thr | 170 | 175 | 180 |
| Gln | Leu | Arg | Val | Leu | Gly | Pro | Glu | Asp | Asp | Leu | Ala | Gly | Met | Phe | 185 | 190 | 195 |
| Leu | Gln | Ile | Phe | Pro | Leu | Ser | Pro | Asp | Pro | Arg | Trp | Gln | Ser | Ser | 200 | 205 | 210 |
| Ser | Pro | Arg | Pro | Val | Ala | Leu | Ala | Leu | Gln | Gln | Ala | Leu | Gly | Gln | 215 | 220 | 225 |
| Glu | Leu | Ala | Arg | Val | Val | Gln | Gly | Ser | Pro | Glu | Val | Pro | Gly | Ile | 230 | 235 | 240 |
| Thr | Val | Arg | Val | Leu | Gln | Ala | Leu | Ala | Thr | Leu | Leu | Ser | Ser | Pro | 245 | 250 | 255 |
| His | Gly | Gly | Ala | Leu | Val | Met | Ser | Met | His | Arg | Ser | His | Phe | Leu | 260 | 265 | 270 |
| Ala | Cys | Pro | Leu | Leu | Arg | Gln | Leu | Cys | Gln | Tyr | Gln | Arg | Cys | Val | 275 | 280 | 285 |
| Pro | Gln | Asp | Thr | Gly | Phe | Ser | Ser | Leu | Phe | Leu | Lys | Val | Leu | Leu | 290 | 295 | 300 |
| Gln | Met | Leu | Gln | Trp | Leu | Asp | Ser | Pro | Gly | Val | Glu | Gly | Gly | Pro | 305 | 310 | 315 |
| Leu | Arg | Ala | Gln | Leu | Arg | Met | Leu | Ala | Ser | Gln | Ala | Ser | Ala | Gly | 320 | 325 | 330 |
| Arg | Arg | Leu | Ser | Asp | Val | Arg | Gly | Gly | Leu | Leu | Arg | Leu | Ala | Glu | 335 | 340 | 345 |
| Ala | Leu | Ala | Phe | Arg | Gln | Asp | Leu | Glu | Val | Val | Ser | Ser | Thr | Val | 350 | 355 | 360 |
| Arg | Ala | Val | Ile | Ala | Thr | Leu | Arg | Ser | Gly | Glu | Gln | Cys | Ser | Val | 365 | 370 | 375 |
| Glu | Pro | Asp | Leu | Ile | Ser | Lys | Val | Leu | Gln | Gly | Leu | Ile | Glu | Val | 380 | 385 | 390 |
| Arg | Ser | Pro | His | Leu | Glu | Glu | Leu | Leu | Thr | Ala | Phe | Phe | Ser | Ala | | | |

| | | | | | | | | | | | | | | |
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| | | | | 395 | | | | | 400 | | | | | 405 |
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| Gly | Pro | Ser | Ser | Gly 455 | Leu | Leu | Val | Asp | Trp 460 | Leu | Glu | Met | Leu | Asp 465 |
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| Thr | Leu | His | Gln | Cys 515 | Ile | Arg | Val | Leu | Leu 520 | Gly | Lys | Ser | Arg | Glu 525 |
| Gln | Arg | Phe | Asp | Pro 530 | Ser | Ala | Ser | Leu | Asp 535 | Phe | Leu | Trp | Ala | Cys 540 |
| Ile | His | Val | Pro | Arg 545 | Ile | Trp | Gln | Gly | Arg 550 | Asp | Gln | Arg | Thr | Pro 555 |
| Gln | Lys | Arg | Arg | Glu 560 | Glu | Leu | Val | Leu | Arg 565 | Val | Gln | Gly | Pro | Glu 570 |
| Leu | Ile | Ser | Leu | Val 575 | Glu | Leu | Ile | Leu | Ala 580 | Glu | Ala | Glu | Thr | Arg 585 |
| Ser | Gln | Asp | Gly | Asp 590 | Thr | Ala | Ala | Cys | Ser 595 | Leu | Ile | Gln | Ala | Arg 600 |
| Leu | Pro | Leu | Leu | Leu 605 | Ser | Cys | Cys | Cys | Gly 610 | Asp | Asp | Glu | Ser | Val 615 |
| Arg | Lys | Val | Thr | Glu 620 | His | Leu | Ser | Gly | Cys 625 | Ile | Gln | Gln | Trp | Gly 630 |
| Asp | Ser | Val | Leu | Gly 635 | Arg | Arg | Cys | Arg | Asp 640 | Leu | Leu | Leu | Gln | Leu 645 |
| Tyr | Leu | Gln | Arg | Pro 650 | Glu | Leu | Arg | Val | Pro 655 | Val | Pro | Glu | Val | Leu 660 |
| Leu | His | Ser | Glu | Gly 665 | Ala | Ala | Ser | Ser | Ser 670 | Val | Cys | Lys | Leu | Asp 675 |
| Gly | Leu | Ile | His | Arg 680 | Phe | Ile | Thr | Leu | Leu 685 | Ala | Asp | Thr | Ser | Asp 690 |

| | | | | | | | | | | | | | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| Ser | Arg | Ala | Leu | Glu | Asn | Arg | Gly | Ala | Asp | Ala | Ser | Met | Ala | Cys | 695 | 700 | 705 |
| Arg | Lys | Leu | Ala | Val | Ala | His | Pro | Leu | Leu | Leu | Leu | Arg | His | Leu | 710 | 715 | 720 |
| Pro | Met | Ile | Ala | Ala | Leu | Leu | His | Gly | Arg | Thr | His | Leu | Asn | Phe | 725 | 730 | 735 |
| Gln | Glu | Phe | Arg | Gln | Gln | Asn | His | Leu | Ser | Cys | Phe | Leu | His | Val | 740 | 745 | 750 |
| Leu | Gly | Leu | Leu | Glu | Leu | Leu | Gln | Pro | His | Val | Phe | Arg | Ser | Glu | 755 | 760 | 765 |
| His | Gln | Gly | Ala | Leu | Trp | Asp | Cys | Leu | Leu | Ser | Phe | Ile | Arg | Leu | 770 | 775 | 780 |
| Leu | Leu | Asn | Tyr | Arg | Lys | Ser | Ser | Arg | His | Leu | Ala | Ala | Phe | Ile | 785 | 790 | 795 |
| Asn | Lys | Phe | Val | Gln | Phe | Ile | His | Lys | Tyr | Ile | Thr | Tyr | Asn | Ala | 800 | 805 | 810 |
| Pro | Ala | Ala | Ile | Ser | Phe | Leu | Gln | Lys | His | Ala | Asp | Pro | Leu | His | 815 | 820 | 825 |
| Asp | Leu | Ser | Phe | Asp | Asn | Ser | Asp | Leu | Val | Met | Leu | Lys | Ser | Leu | 830 | 835 | 840 |
| Leu | Ala | Gly | Leu | Ser | Leu | Pro | Ser | Arg | Asp | Asp | Arg | Thr | Asp | Arg | 845 | 850 | 855 |
| Gly | Leu | Asp | Glu | Glu | Gly | Glu | Glu | Glu | Ser | Ser | Ala | Gly | Ser | Leu | 860 | 865 | 870 |
| Pro | Leu | Val | Ser | Val | Ser | Leu | Phe | Thr | Pro | Leu | Thr | Ala | Ala | Glu | 875 | 880 | 885 |
| Met | Ala | Pro | Tyr | Met | Lys | Arg | Leu | Ser | Arg | Gly | Gln | Thr | Val | Glu | 890 | 895 | 900 |
| Asp | Leu | Leu | Glu | Val | Leu | Ser | Asp | Ile | Asp | Glu | Met | Ser | Arg | Arg | 905 | 910 | 915 |
| Arg | Pro | Glu | Ile | Leu | Ser | Phe | Phe | Ser | Thr | Asn | Leu | Gln | Arg | Leu | 920 | 925 | 930 |
| Met | Ser | Ser | Ala | Glu | Glu | Cys | Cys | Arg | Asn | Leu | Ala | Phe | Ser | Leu | 935 | 940 | 945 |
| Ala | Leu | Arg | Ser | Met | Gln | Asn | Ser | Pro | Ser | Ile | Ala | Ala | Ala | Phe | 950 | 955 | 960 |
| Leu | Pro | Thr | Phe | Met | Tyr | Cys | Leu | Gly | Ser | Gln | Asp | Phe | Glu | Val | 965 | 970 | 975 |
| Val | Gln | Thr | Ala | Leu | Arg | Asn | Leu | Pro | Glu | Tyr | Ala | Leu | Leu | Cys | | | |

| | | | |
|---|-----|-----|-----|
| Gln Lys Gly Asp Val Val Asp Val Tyr Gln Arg Glu Phe Leu Ala | 35 | 40 | 45 |
| Leu Arg Asp Arg Leu His Ala Ala Glu Gln Glu Ser Leu Lys Arg | 50 | 55 | 60 |
| Ser Lys Glu Leu Asn Leu Val Leu Asp Glu Ile Lys Arg Ala Val | 65 | 70 | 75 |
| Ser Glu Arg Gln Ala Leu Arg Asp Gly Asp Gly Asn Arg Thr Trp | 80 | 85 | 90 |
| Gly Arg Leu Thr Glu Asp Pro Arg Leu Lys Pro Trp Asn Gly Ser | 95 | 100 | 105 |
| His Arg His Val Leu His Leu Pro Thr Val Phe His His Leu Pro | 110 | 115 | 120 |
| His Leu Leu Ala Lys Glu Ser Ser Leu Gln Pro Ala Val Arg Val | 125 | 130 | 135 |
| Gly Gln Gly Arg Thr Gly Val Ser Val Val Met Gly Ile Pro Ser | 140 | 145 | 150 |
| Val Arg Arg Glu Val His Ser Tyr Leu Thr Asp Thr Leu His Ser | 155 | 160 | 165 |
| Leu Ile Ser Glu Leu Ser Pro Gln Glu Lys Glu Asp Ser Val Ile | 170 | 175 | 180 |
| Val Val Leu Ile Ala Glu Thr Asp Ser Gln Tyr Thr Ser Ala Val | 185 | 190 | 195 |
| Thr Glu Asn Ile Lys Ala Leu Phe Pro Thr Glu Ile His Ser Gly | 200 | 205 | 210 |
| Leu Leu Glu Val Ile Ser Pro Ser Pro His Phe Tyr Pro Asp Phe | 215 | 220 | 225 |
| Ser Arg Leu Arg Glu Ser Phe Gly Asp Pro Lys Glu Arg Val Arg | 230 | 235 | 240 |
| Trp Arg Thr Lys Gln Asn Leu Asp Tyr Cys Phe Leu Met Met Tyr | 245 | 250 | 255 |
| Ala Gln Ser Lys Gly Ile Tyr Tyr Val Gln Leu Glu Asp Asp Ile | 260 | 265 | 270 |
| Val Ala Lys Pro Asn Tyr Leu Ser Thr Met Lys Asn Phe Ala Leu | 275 | 280 | 285 |
| Gln Gln Pro Ser Glu Asp Trp Met Ile Leu Glu Phe Ser Gln Leu | 290 | 295 | 300 |
| Gly Phe Ile Gly Lys Met Phe Lys Ser Leu Asp Leu Ser Leu Ile | 305 | 310 | 315 |
| Val Glu Phe Ile Leu Met Phe Tyr Arg Asp Lys Pro Ile Asp Trp | | | |

| | | | | | | | | | | | | | | |
|-----|-----|-----|-----|------------|-----|-----|-----|-----|------------|-----|-----|-----|-----|------------|
| | | | | 320 | | | | | 325 | | | | | 330 |
| Leu | Leu | Asp | His | Ile 335 | Leu | Trp | Val | Lys | Val 340 | Cys | Asn | Pro | Glu | Lys 345 |
| Asp | Ala | Lys | His | Cys 350 | Asp | Arg | Gln | Lys | Ala 355 | Asn | Leu | Arg | Ile | Arg 360 |
| Phe | Lys | Pro | Ser | Leu 365 | Phe | Gln | His | Val | Gly 370 | Thr | His | Ser | Ser | Leu 375 |
| Ala | Gly | Lys | Ile | Gln 380 | Lys | Leu | Lys | Asp | Lys 385 | Asp | Phe | Gly | Lys | Gln 390 |
| Ala | Leu | Arg | Lys | Glu 395 | His | Val | Asn | Pro | Pro 400 | Ala | Glu | Val | Ser | Thr 405 |
| Ser | Leu | Lys | Thr | Tyr 410 | Gln | His | Phe | Thr | Leu 415 | Glu | Lys | Ala | Tyr | Leu 420 |
| Arg | Glu | Asp | Phe | Phe 425 | Trp | Ala | Phe | Thr | Pro 430 | Ala | Ala | Gly | Asp | Phe 435 |
| Ile | Arg | Phe | Arg | Phe 440 | Phe | Gln | Pro | Leu | Arg 445 | Leu | Glu | Arg | Phe | Phe 450 |
| Phe | Arg | Ser | Gly | Asn 455 | Ile | Glu | His | Pro | Glu 460 | Asp | Lys | Leu | Phe | Asn 465 |
| Thr | Ser | Val | Glu | Val 470 | Leu | Pro | Phe | Asp | Asn 475 | Pro | Gln | Ser | Asp | Lys 480 |
| Glu | Ala | Leu | Gln | Glu 485 | Gly | Arg | Thr | Ala | Thr 490 | Leu | Arg | Tyr | Pro | Arg 495 |
| Ser | Pro | Asp | Gly | Tyr 500 | Leu | Gln | Ile | Gly | Ser 505 | Phe | Tyr | Lys | Gly | Val 510 |
| Ala | Glu | Gly | Glu | Val 515 | Asp | Pro | Ala | Phe | Gly 520 | Pro | Leu | Glu | Ala | Leu 525 |
| Arg | Leu | Ser | Ile | Gln 530 | Thr | Asp | Ser | Pro | Val 535 | Trp | Val | Ile | Leu | Ser 540 |
| Glu | Ile | Phe | Leu | Lys 545 | Lys | Ala | Asp | | | | | | | |

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